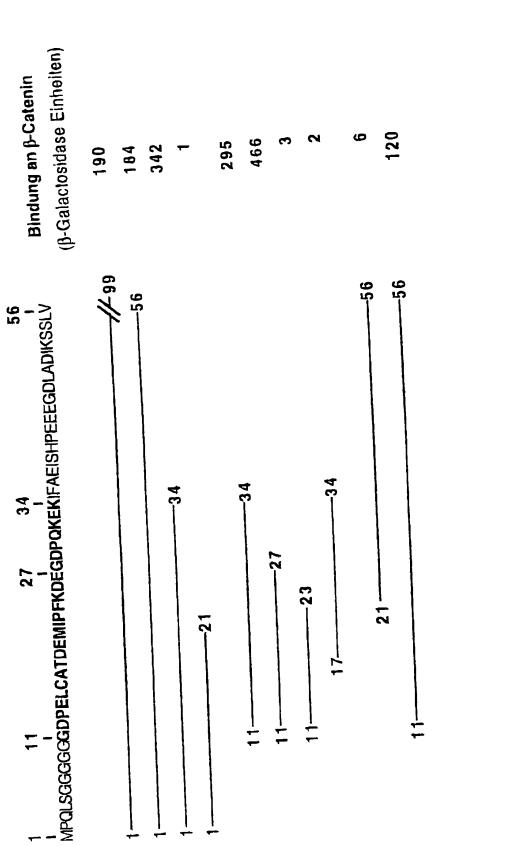
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BERICHTIGTES BLATT (REGEL 91) ISA / EP

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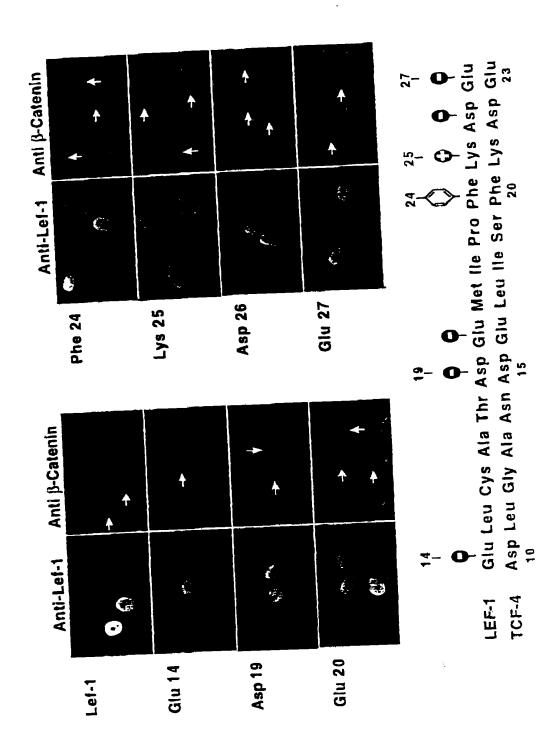


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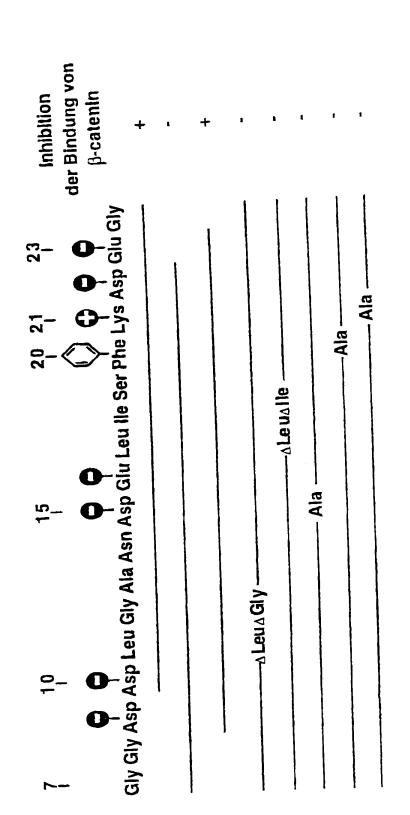
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PCT/DE99/00554

Inhibition der LEF-1/β-catenin Interaktion durch synthetische Peptide aus der Minimalen Bindungs-Domäne

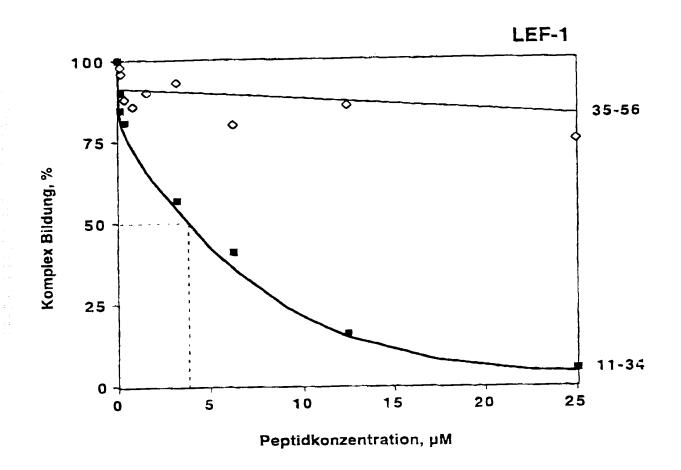
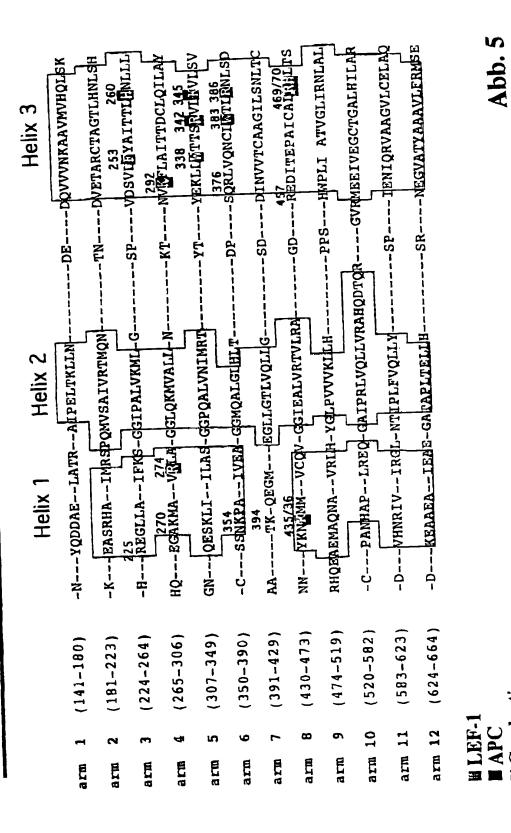


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Conductin E-Cadherin

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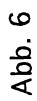


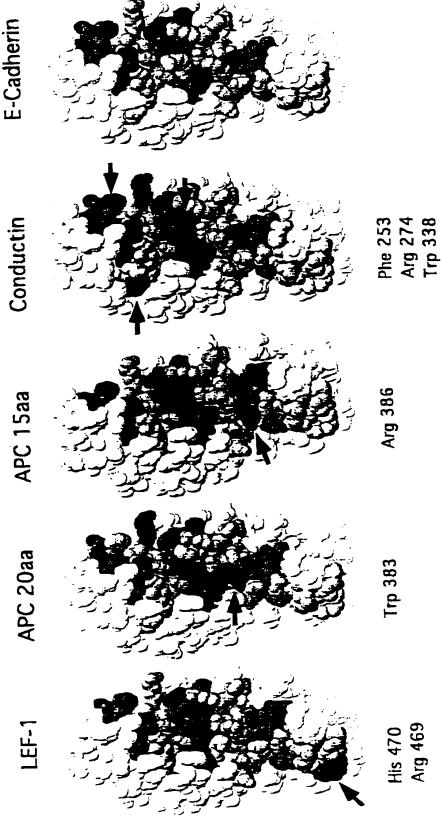
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Interaktion von β -Catenin Mutanten mit:

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Interaktions-Reporteraktivität:

0 - 30 % 30- 60 % ➤ : Mutation für Interaktionspartner spezifisch

BERICHTIGTES BLATT (REGEL 91)
ISA / EP

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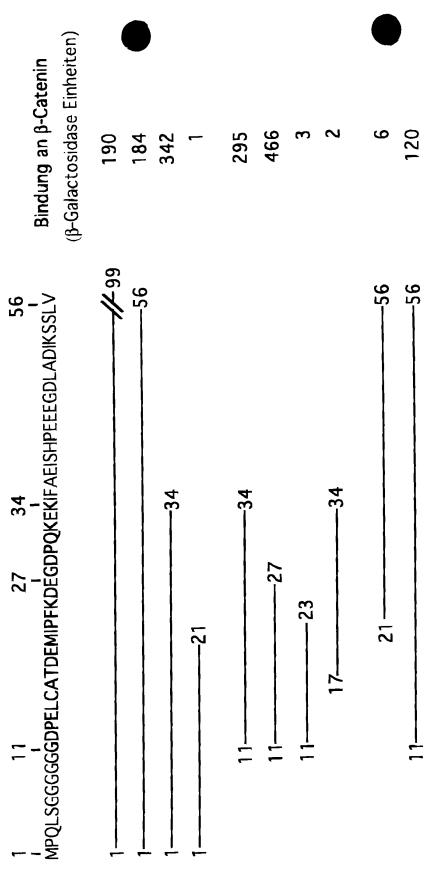


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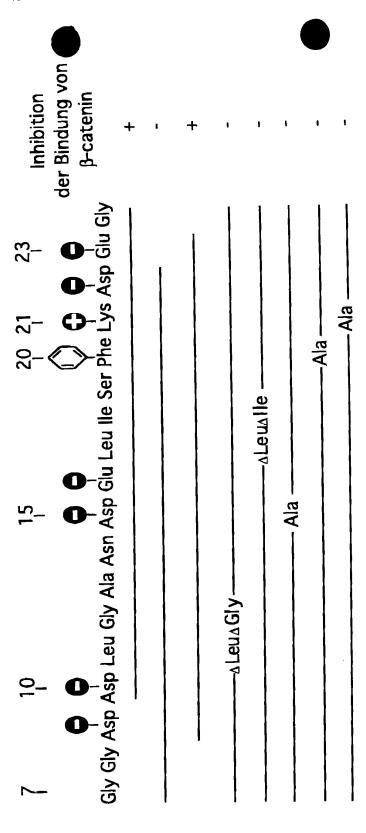


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Inhibition der LEF-1/β-catenin Interaktion durch synthetische Peptide aus der Minimalen Bindungs-Domäne

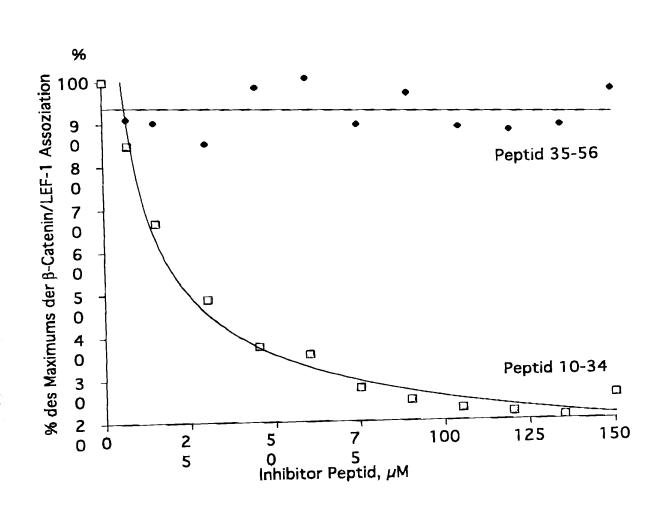


Abb. 3

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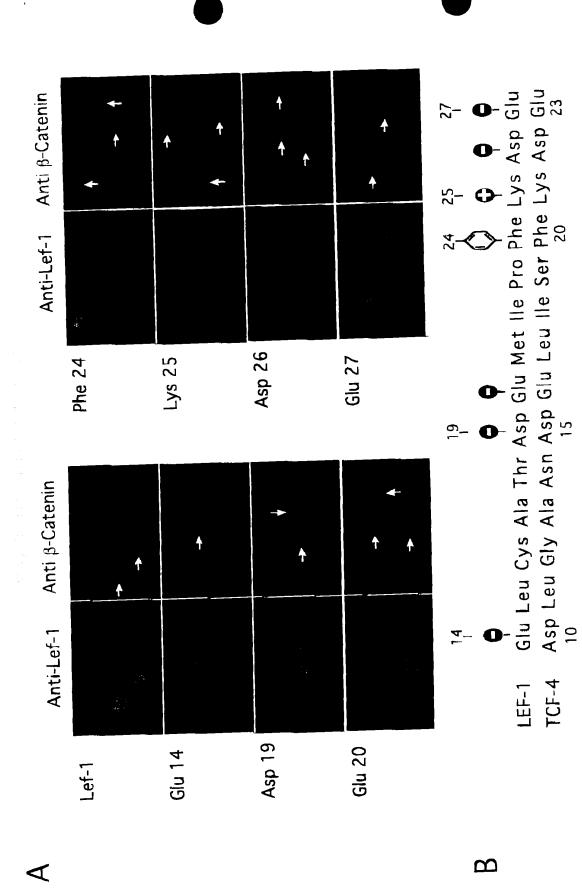
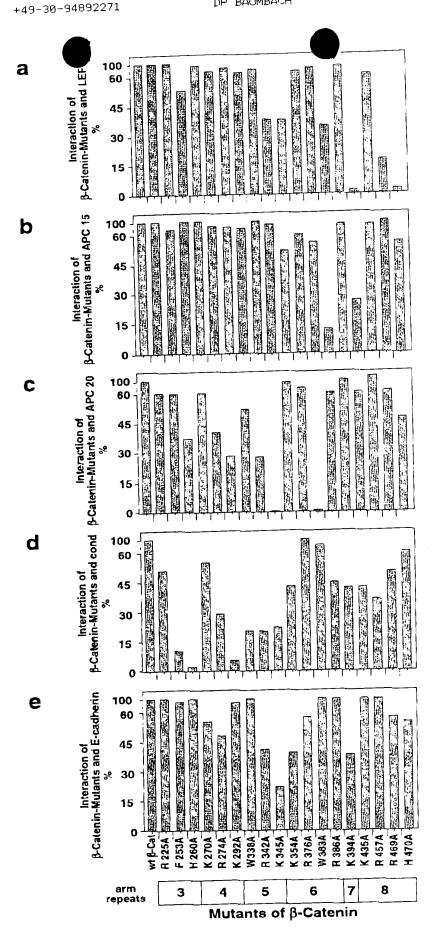
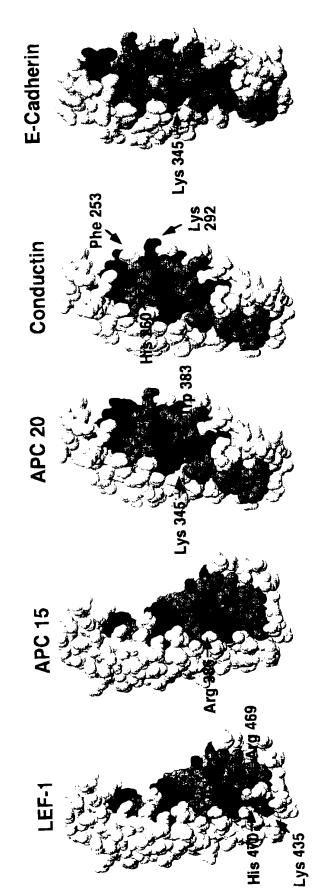


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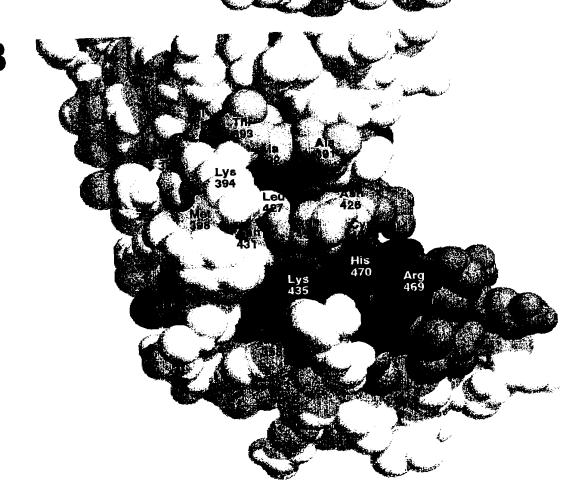
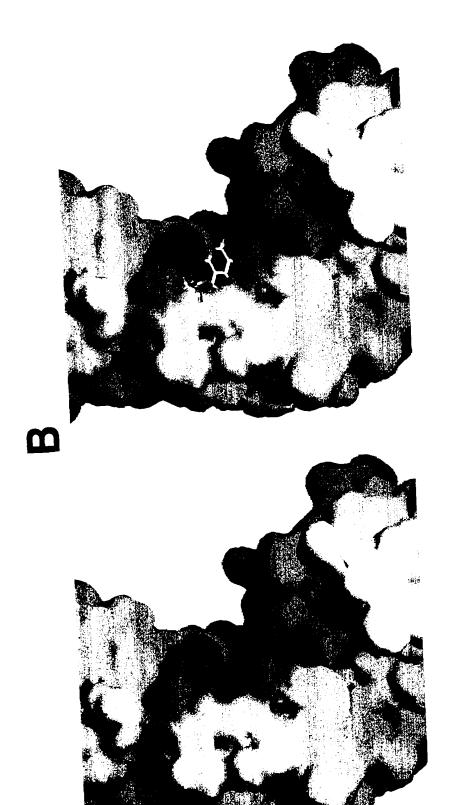
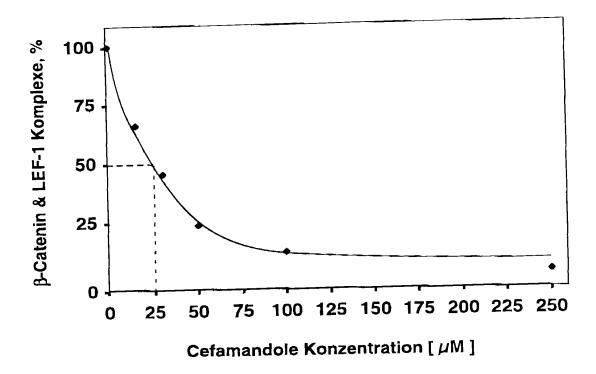


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Aminosäuresequenz des humanen \(\beta\)-Catenin (Armadillo-Repeats 3-9)

+49-30-94892271

arm	m	(224-264)	AREGILATERSGGIPAL VRALG SPVDSVLFYAITTERNALEE
arm	41	(265-306)	HQEGA MAVRLAGGLQKMVALLNKTNVKFLAITTBCLQILAY
arm	Ŋ	(307-349)	GNQESKLIILASGGPQALVNIMRTYTYEKKATWTTSRVLKVLSV
arm	9	(350-390)	CSSNKPAIVEAGGMQALGLHLTDFSQRLVQNCLWTLRNLSD
arm	7	(391-429)	AATKQEGMEGLLGTLWOLLGSDDINVVTCAAGILSNLTC
arm	æ	(430-473)	NNY KNKMMYEQVGGIEALVRTVLRAGDREDITEPAICALRHLTS
arm	6	arm 9 (474-519)	AHOEAEMHONAUREHYGLPUVUYKLLHPPSHWPLFKATUGLFRUEAL

Tab.2 Interaktion von β-Catenin Mutanten mit LEF-1, APC (20 und 15 Aminosäure -Repeats), Conductin und E-Cadherin

		Interaktion mit				
β-Catenin Mutanten	arm. Einh.	LEF-1	APC-20	APC-15	Conductin	E-Cadherin
Phe 253	3	-	40	-	17	-
His 260	3	50	40	100	10	100
Arg 274	4	-	40	-	29	50
Lys 292	4	_	28	-	5	-
Trp 338	5	_	55	_	20	-
Arg 342	5	-	29	-	20	41
Lys 345	5	38	0	-	22	27
Lys 354	6	38	-	54	43	40
Trp 383	6	_	0	59	_	-
Arg 386	6	35		12	45	-
Lys 394	7	_	_	-	42	-
Lys 435	8	-	_	30	42	-
Arg 457	8	-	-	-	36	-
Arg 469	8	17	-	-	-	50
His 470	8	2	47	60	-	_

Die Werte geben den prozentualen Anteil der jeweiligen Interaktion im Vergleich zu der mit Wildtyp-β-Catenin an. Durch "-" gekennzeichnete Interaktionen entsprechen, 60 -100 % der Wildtyp-Interaktion.

s.

Tab. 3





MOLEKÜLSTRUKTUR	Substanzname	MDL-Nr.
- HO OH	(-)-ESEROLINE FUMARATE	MFCD00055202
- 6 - · · · · · · · · · · · · · · · · ·		
H.C C		
₩		
Eri,		
OH Chiral	N-ACETYL-MURAMYL-ALA-ISOGLN-OH	MFCD00065478
HO. N. CH,	N-ACETTE-MONAWITE-ADA-ISOGEN-OIT	WFCD00003478
o CHEH, O		
- SON TOTAL		
но		
OH	3,6-DIHYDROXYBENZONORBORNANE	MFCD00077441
- OH H		
Chiral Chiral	N-ACETYL-MURAMYL-ALA-D-ISOGLN-OH	MFCD00077638
_ о		
CH ₃ N ₄ OH —		
OH CH, NO CHANN		
OH OF NH		
Chirel	(-)-COTININE	MFCD00077696
N N		
- ĊH ₃ -		
	CEFAMANDOLE SODIUM SALT	MECDOOCCO
	CLFAIVAINDOLE SUDIUIVI SALI	MFCD00082385
N S		
ONN		
0 N-N		
, <u>-</u>		



D D O OH OH	(+/-)-NICOTINE-D3 SALICYLATE SALT	MFCD00083448
	BENZYL 1,2-DIPHENYL-4-HYDROXY-5-OXO-C PYRROLINE-3-CARBOXYLATE	3-MFCD00088051
	1-(4-FLUOROPHENYL)-3- PHENYLPYRROLIDINE-2,5-DIONE	MFCD00097831
	3-PHENYL-1-(2- THIENYLMETHYL)PYRROLIDINE-2,5-DIONE	MFCD00097832
F N O F F F	N1-(4-[2,5-DIOXO-1-[4- (TRIFLUOROMETHOXY)PHENYLJTETRAHYDR O-1H-PYRROL-3-YLJPHENYL)-2,2,2- TRIFLUOROACETAMIDE	MFCD00100474
	1,3-DIPHENYLCYCLOPENTANE-2,4,5-TRIONE	MFCD00101320





	ETHYL 2-OXO-2-[(2-OXOAZEPAN-3- YL)AMINO]ACETATE	MFCD00103142
	3-BENZOYL-1,2,3,10B-TETRAHYDRO- PYRROLO(2,1-A)ISOQUINOLINE-1- CARBONITRILE	MFCD00123443
HO Chiral _	L-TRANS-EPOXYSUCCINYL-LEU-3- METHYLBUTYLAMIDE	MFCD00132882
N _{1,1} CH ₃ CH ₃		
CH ₃		
HO Catest 2 Autol	AJMALINE	MFCD00135652
N, OH		\
CH ³ H CH,		,
Chirel	(2S,3R)-3-PHENYLPYRROLIDINE-2- CARBOXYLIC ACID	MFCD00142984
		,
N OH		
ОНОН	4-(2-HYDROXYOCTAHYDRO-1,3,4-METHENO 2H-CYCLOBUTA(CD)PENTALEN-2- YL)PHENOL	MFCD00155174





	METHYL C-4-(4-METHOXYPHENYL)-2- BENZYL-7-PHENYL-6,8-DIOXO-3,7- DIAZABICYCLO[3.3.0]-OCTANE-R-2- CARBOXYLATE	MFCD00202518
	METHYL C-4-(4-METHOXYPHENYL)-2- BENZYL-7-(4-CHLOROPHENYL)-6,8-DIOXO- 3,7-DIAZABICYCLO[3,3.0]-OCTANE-R-2- CARBOXYLATE	MFCD00202519
CI-O-CH;		
CIH N H	(+/-)-EPIBATIDINE DIHYDROCHLORIDE	MFCD00210196
CIH ————————————————————————————————————	4-CYCLOPENTYL-NAPHTHALEN-1-YLAMINE HYDROCHLORIDE	MFCD00227852
СН,	5-(2,3-DIMETHOXY-PHENYL)-PYRAZOLIDIN- 3-ONE	MFCD00228403
H ₃ C	5-(3,4-DIMETHOXY-PHENYL)-PYRAZOLIDIN 3-ONE	- MFCD00229211
O CH ₃		





0 0-N ^t	5-CYCLOPENTYL-2-METHOXY-4-NITRO- PHENOL	MFCD00230901
CH,		
OH Chuşi	AC-(6-O-STEAROYL)-MURAMYL-ALA-D- ISOGLUTAMINE	MFCD00236777
H _C C O		
	ANTHO-RNAMIDE	MFCD00236789
CH ₃ CHVa.	1,3,5(10),6,8(14BETA)-ESTRAPENTAEN-3- OL-17-ONE ACETATE	MFCD00271642
H ₀ C O F	NORFLUOROCURARINE	MFCD00274483
N N O	4,5-DIPHENYLPYRAZOLIDIN-3-ONE	MFCD00277796
C-**•	4-(4-METHOXY-PHENYL)-2-OXO- PYRROLIDINE-3-CARBOXYLIC ACID	MFCD00297824
СН		





CIA	(+)-EPIBATIDINE HYDROCHLORIDE	MFCD00467208
H	FLUOROCURARINE CHLORIDE	MFCD00467712
	PLOUROCUPANINE CALORIDE	WFCD00467712
CH ₃ Crites		
— A (GH -		
		j
т сн,		
0	(1'S,2'S)-NICOTINE 1'-OXIDE	MFCD00869528
WATEN	(13,23) INICOTINE 1-OXIDE	MECDOOS69528
N-O-		
—		
	SPECS	MFCD01114864
CIH	SPECS CIF7952	1
CI I	2-ETHYL-2-(3-	MFCD01314146
CH ₃	METHOXYPHENYL)PYRROLIDINE	<u> </u>
O CH3		
· · · · · · · · · · · · · · · · · · ·		
CH,	N-(TERT-BUTYL)-2-(ISOXAZOL-5-	MFCD01314517
N CH,	YLCARBONYL)DECAHYDROISOQUINOLINE-3-	
	CARBOXAMIDE	
		<u> </u>

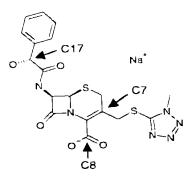
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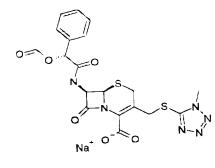


Positiv-Liste



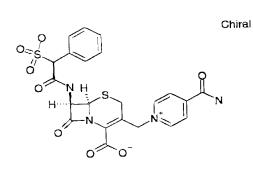
Molekül-Klasse IA (Inhibitoren vom Cefamandole-Typ)

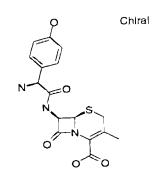




Cefamandole

Cefamandole-Nafate

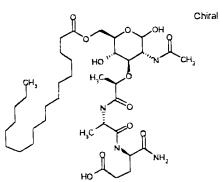




Cefsulodin

Cefadroxil

Molekül-Klasse IB



AC-(6-O-STEAROYL)-MURAMYL-ALA-D-ISOGLUTAMINE



Positiv-Liste



Molekül-Klasse IC

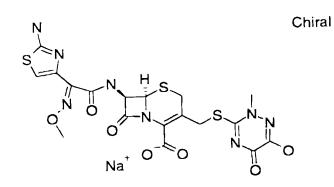
3,6-DIHYDROXYBENZONORBORNANE

Moleküle mit Modulationswirkung für Molekülklasse I

1-(4-FLUOROPHENYL)-3-PHENYLPYRROLIDINE-2,5-DIONE

(-)-COTININE

4-(2-HYDROXYOCTAHYDRO-1,3,4-METHENO-2H-CYCLOBUTA(CD)PENTALEN-2-YL)PHENOL Positiv-Liste



Cefatriaxone

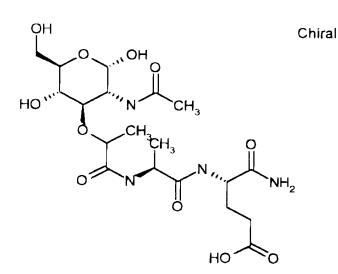
L-TRANS-EPOXYSUCCINYL-LEU-3-METHYLBUTYLAMIDE

ANTHO-RNAMIDE

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Positiv-Liste



N-ACETYL-MURAMYL-ALA-ISOGLN-OH

1,3,5(10),6,8(14BETA)-ESTRAPENTAEN-3-OL-17-ONE ACETATE